

IN THE SPECIFICATION:

Please amend the specification as shown:

Please delete paragraph [019] on page 6, and replace it with the following paragraph:

[019] **FIG. 1.** Comparison of deduced *D. melanogaster* cDNA SD07655 (SEQ ID NO: 1) and human MRP1 (SEQ ID NO: 6) amino acid sequences. The two amino acid sequences were aligned using ClustalW. Identical residues are marked with shading. The transmembrane regions are noted by a fine underline and the ATP-binding domains are noted by a bold underline. The amino acids derived from exons 4 and 8 of the dMRP gene are presented in bold characters. The small vertical lines above and below the amino acids denote the exon junctions with the type of splice junction marked by a number noting the class: 0, 1 or 2. The dMRP amino acid sequence differs from that of sequence AY069827 at the following positions : L/V pos. 124, M/L pos. 318 and I/T pos. 448.

Please delete paragraph [022] on pages 7-8, and replace it with the following paragraph:

[022] **FIG. 4.** Amino acid alignment of dMRP variable exon 4 (A) (SEQ ID NOS 7 & 8) and 8 (B) (SEQ ID NOS 9-15) encoded peptides with the cognate peptides from other organisms. The variant dMRP peptide sequence and the equivalent sequences

from *Drosophila* sulfonylurea receptor (Dsur, NG_000795) (SEQ ID NOS 134 & 138) and three human MRPs (MRP1, NM_004996 (SEQ ID NOS 131 & 135); MRP2, NP_005836 (SEQ ID NOS 132 & 136); and MRP3, Y17151 (SEQ ID NOS 133 & 137)) were aligned using ClustalW. Pfam (SEQ ID NO: 139) refers to pfam00664, a consensus sequence for ABC transporter Membrane Spanning Domains. Gaps were introduced to maximize sequence identity and are shown by a horizontal dash. Residues that are identical in at least half of the sequences have their background shaded and those present in more than half of the sequences are listed in the consensus (Cons). (C) Dendrogram constructed with the data of part (B) of the Figure (see *infra* for details).

Please delete paragraph [024] on page 8, and replace it with the following paragraph:

[024] **Fig. 6.** Comparison of deduced *A. gambiae* gMRP1a-d (SEQ ID NOS 2-5), *Drosophila melanogaster* dMRP (SEQ ID NO: 1), and human MRP1 (SEQ ID NO: 6) amino acid sequences. The alignment was produced using ClustalW. Identical residues in at least half of the sequences are marked with shading. The different topological regions are indicated in bold and italic above the sequences, and are delimitated by vertical bars. *MSD1-3*, Membrane Spanning Domains 1 to 3; *L₀*, cytoplasmic loop; *NBD1-2*, Nucleotide Binding Domain, *Linker*, region linking the two halves of the protein. Walker A and Walker B are indicated as *A* and *B*, and their sequences are

marked in bold, as well as the signature (C) of ABC transporters. The vertical lines in bold inside the amino acid sequences denote the exon junctions. Where several genes shared the same site, this one was emphasized by a delimitating box.

Please delete paragraph [052] on pages 21-22, and replace it with the following paragraph:

[052] DNA (10 µg) was digested with either *Bam*H or *Hind*III and the fragments were separated by electrophoresis on a 0.8% agarose gel. Following transfer to Hybond-N nylon membrane and fixation, hybridization was carried out at 65°C (in 1% BSA, 0.25 M NaH₂PO₄ pH 7.2, 1 mM EDTA, 150 µg/ml salmon sperm DNA) with a PCR-derived *dMRP* probe covering 378 bases (forward primer:

GATCCGTTATTCCCTTGCCGC (**SEQ ID NO: 53**); reverse primer:
TCCAGGGCAGTGATTACCACT (**SEQ ID NO: 54**). After hybridization, the blot was washed (in 40 mM NaH₂PO₄ pH 7.2, 1% SDS, and 1 mM EDTA) 1X at RT and 2X at 65 C°.

Please delete Table 3, on page 31, and replace it with the Table at Tab A.

Please delete Table 5, on page 39, and replace it with the Table at Tab B.



TABLE 3. Intron-exon organization of the *Drosophila dMRP* gene

Exon	3' acceptora (SEQ ID NOS 55- n° Size (bp) 72, respectively, in exon locationb order of appearance)	5' donor (SEQ ID NOS 73- 90, respectively, in order of appearance)	Intron		
			size (bp)	n° Phase	Size (bp)
1	181	-127•54	TTCTGG / gtgagt	1	0
2	1512	129•1640	ATTAAG /gtgagt	2	0
3	138	1776•1913	TTCCCTG / gtaaga	3	0
4a	147	2042•2188	GCCGAG / gtacag	4	0
4b	147	2335•2481	GTGCAC / gtaagt	5	0
5	85	3282•3366	CTAAAC / gtaaga	6	1
6	820	3429•4248	TTCCAT / gtaagt	7	2
7	371	4316•4686	GCCAAG / gtaagt	8	1
8a	221	5591•5811	TATATG / gtaatt	9	0
8b	221	6148•6368	TTTGCG / gtaatt	10	0
8c	221	6754•6974	TTTGCG / gtaaat	11	0
8d	221	7500•7720	TTTCGGG / gtaaag	12	0
8e	224	8412•8635	TTTATG / gttattt	13	0
8f	221	13605•13825	TTTCAG / gtaatc	14	0
8g	221	14967•15187	TTCGAG / gtaatt	15	0
9	218	15528•15745	AGATCG / gtaatgt	16	2
10	507	15810•16316	GTTCAG / gtaagc	17	2
11	382	16376•16757	ATTCAAG / gttgggt	18	0
12	393	21549•21941			4791

Table 5. Organization of exon-intron junctions in the gMRPs

Name	Location on protein sequence ^a	Size (bp)	Exon			Intron		
			3' acceptor ^b <u>(SEQ ID NOS</u>		5' donor ^b <u>(SEQ ID NOS</u>	Name	Phase	Size (bp)
			<u>91-110</u>	respectively, in order of appearance)	<u>111-130</u> , respectively, in order of appearance)			
<i>MRP1a</i>	1	165	CCCTTG/gtgaga	1	0	83		
	2	234	TCCCTG/gtaagg	2	0	202		
	3	566	GCTGAG/gtaagt	3	2	224		
	4	3638						
<i>MRP1b</i>	1	ND ^c						
	2	13	TTTTGG/gtaagt	1	0	603		
	3	300	GCTTAT/gtaagt	2	0	76		
	4	2144	ATACCA/gtaagt	3	2	63		
	5	1458	CTTCAG/gtatgt	4	2	73		
	6	382	ATTCAg/gtaaga	5	0	65		
<i>MRP1c</i>	1	493						
	2	418	GCTTAT/gtgagt	1	0	69		
	3	662	GATGCAG/gtaagt	2	2	96		
	4	113	TTATCA/gtaagt	3	2	61		
	5	334	ATGAAG/gtaagt	4	1	60		
	6	833	CTTCAG/gtttagt	5	2	63		
	7	858	ATTCAg/gtgaga	6	0	71		
	8	1315						
	9	1442						
<i>MRP1d</i>	1	ND ^c						
	2	113						
	3	662	GCTTAT/gtgagt	1	0	69		
	4	1497	CATGCCA/gtacgt	2	2	110		
	5	77	ATACCA/gtaagt	3	2	65		
	6	1369	AAGACGG/ttaggt	4	1	98		
	7	382	CTTCAG/gtatct	5	2	73		
	8	293	ATTCAg/gtaaga	6	0	65		
	9	ND ^c						
	10	113						
	11	334						
	12	833						
	13	859						
	14	1317						
	15	1444						

a) The numbering is based on amino acid one being the putative first Met.

b) Capital letters are used for the sequence in the exon and small case letters for sequence in the intron.

c) Not Determined.

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b) Capital letters are used for the sequence in the exon and small case letters for sequence in the intron.

c) Not Determined.